SEQUENCE LISTING

- <110> MacKinnon, Roderick The Rockefeller University
- <120> Assays for Screening Compounds Which Interact With Cation Channel Proteins, Mutant Prokaryotic Cation Channel Proteins, and Uses Thereof
- <130> 018512-002901US
- <140> US 09/275,252
- <141> 1999-03-24
- <150> US 09/045,529
- <151> 1998-03-20
- <150> US 09/054,347
- <151> 1998-04-02.
- <150> WO PCT/US99/06307
- <151> 1999-03-22
- <160> 42
- <170> PatentIn Ver. 2.1
- <210> 1
- <211> 160
- <212> PRT
- <213> Streptomyces lividans
- <400> 1
- Met Pro Pro Met Leu Ser Gly Leu Leu Ala Arg Leu Val Lys Leu Leu 1 5 10 15
- Leu Gly Arg His Gly Ser Ala Leu His Trp Arg Ala Ala Gly Ala Ala 20 25 30
- Thr Val Leu Leu Val Ile Val Leu Leu Ala Gly Ser Tyr Leu Ala Val
- Leu Ala Glu Arg Gly Ala Pro Gly Ala Gln Leu Ile Thr Tyr Pro Arg
 50 55 60
- Ala Leu Trp Trp Ser Val Glu Thr Ala Thr Thr Val Gly Tyr Gly Asp
 65 70 75 80
- Leu Tyr Pro Val Thr Leu Trp Gly Arg Leu Val Ala Val Val Met
 85 90 95
- Val Ala Gly Ile Thr Ser Phe Gly Leu Val Thr Ala Ala Leu Ala Thr 100 105 110
- Trp Phe Val Gly Arg Glu Gln Glu Arg Arg Gly His Phe Val Arg His
 115 120 125
- Ser Glu Lys Ala Ala Glu Glu Ala Tyr Thr Arg Thr Thr Arg Ala Leu 130 135 140

His Glu Arg Phe Asp Arg Leu Glu Arg Met Leu Asp Asp Asn Arg Arg 145 150 155 160

<210> 2

<211> 417

<212> PRT

<213> Escherichia coli

<400> 2

Met Ser His Trp Thr Thr Phe Lys Gln Thr Ala Thr Lys Leu Trp Val

Thr Leu Arg His Asp Ile Leu Ala Leu Ala Val Phe Leu Asn Gly Leu 20 25 30

Leu Ile Phe Lys Thr Ile Tyr Gly Met Ser Val Asn Leu Leu Asp Ile 35 40 45

Phe His Ile Lys Ala Phe Ser Glu Leu Asp Leu Ser Leu Leu Ala Asn 50 55 60

Ala Pro Leu Phe Met Leu Gly Val Phe Leu Val Leu Asn Ser Ile Gly 65 70 75 80

Leu Leu Phe Arg Ala Lys Leu Ala Trp Ala Ile Ser Ile Ile Leu Leu 85 90 95

Leu Ile Ala Leu Ile Tyr Thr Leu His Phe Tyr Pro Trp Leu Lys Phe
100 105 110

Ser Ile Gly Phe Cys Ile Phe Thr Leu Val Phe Leu Leu Ile Leu Arg 115 120 125

Lys Asp Phe Ser His Ser Ser Ala Ala Ala Gly Thr Ile Phe Ala Phe 130 135 140

Ile Ser Phe Thr Thr Leu Leu Phe Tyr Ser Thr Tyr Gly Ala Leu Tyr 145 150 155 160

Leu Ser Glu Gly Phe Asn Pro Arg Ile Glu Ser Leu Met Thr Ala Phe 165 170 175

Tyr Phe Ser Ile Glu Thr Met Ser Thr Val Gly Tyr Gly Asp Ile Val 180 185 190

Pro Val Ser Glu Ser Ala Arg Leu Phe Thr Ile Ser Val Ile Ile Ser 195 200 205

Gly Ile Thr Val Phe Ala Thr Ser Met Thr Ser Ile Phe Gly Pro Leu 210 215 220

Ile Arg Gly Gly Phe Asn Lys Leu Val Lys Gly Asn Asn His Thr Met 225 230 235 240

His Arg Lys Asp His Phe Ile Val Cys Gly His Ser Ile Leu Ala Ile 245 250 255

Asn Thr Ile Leu Gln Leu Asn Gln Arg Gly Gln Asn Val Thr Val Ile 260 265 270 Ser Asn Leu Pro Glu Asp Asp Ile Lys Gln Leu Glu Gln Arg Leu Gly 275 280 285

Asp Asn Ala Asp Val Ile Pro Gly Asp Ser Asn Asp Ser Ser Val Leu 290 295 300

Lys Lys Ala Gly Ile Asp Arg Cys Arg Ala Ile Leu Ala Leu Ser Asp 305 310 315 320

Asn Asp Ala Asp Asn Ala Phe Val Val Leu Ser Ala Lys Asp Met Ser 325 330 335

Ser Asp Val Lys Thr Val Leu Ala Val Ser Asp Ser Lys Asn Leu Asn 340 345 350

Lys Ile Lys Met Val His Pro Asp Ile Ile Leu Ser Pro Gln Leu Phe 355 360 365

Gly Ser Glu Ile Leu Ala Arg Val Leu Asn Gly Glu Glu Ile Asn Asn 370 380

Asp Met Leu Val Ser Met Leu Leu Asn Ser Gly His Gly Ile Phe Ser 385 390 395 400

Asp Asn Asp Glu Gln Glu Thr Lys Ala Asp Ser Lys Glu Ser Ala Gln 405 410 415

Lys

<210> 3

<211> 58

<212> PRT

<213> Clostridium acetobutylicum

<400> 3

Ser Leu Gly Asn Ala Leu Trp Trp Ser Phe Val Thr Ile Thr Thr Val 1 5 10 15

Gly Tyr Gly Asp Ile Ser Pro Ser Thr Pro Phe Gly Arg Val Ile Ala 20 25 30

Ser Ile Leu Met Leu Ile Gly Ile Gly Phe Leu Ser Met Leu Thr Gly
35 40 45

Thr Ile Ser Thr Phe Phe Ile Ser Lys Lys
50 55

<210> 4

<211> 616

<212> PRT

<213> Drosophila melanogaster

<400> 4

Met Ala Ala Val Ala Gly Leu Tyr Gly Leu Gly Glu Asp Arg Gln His
1 5 10 15

Arg Lys Lys Gln Gln Gln Gln Gln His Gln Lys Glu Gln Leu Glu
20 25 30

Gln Lys Glu Glu Gln Lys Lys Ile Ala Glu Arg Lys Leu Gln Leu Arg 35 40 45 and the same of th

Glu Gln Gln Leu Gln Arg Asn Ser Leu Asp Gly Tyr Gly Ser Leu Pro 50 55 60

Lys Leu Ser Ser Gln Asp Glu Glu Gly Gly Ala Gly His Gly Phe Gly 65 70 75 80

Gly Gly Pro Gln His Phe Glu Pro Ile Pro His Asp His Asp Phe Cys
85 90 95

Glu Arg Val Val Ile Asn Val Ser Gly Leu Arg Phe Glu Thr Gln Leu
100 105 110

Arg Thr Leu Asn Gln Phe Pro Asp Thr Leu Leu Gly Asp Pro Ala Arg 115 120 125

Arg Leu Arg Tyr Phe Asp Pro Leu Arg Asn Glu Tyr Phe Phe Asp Arg 130 135 140

Ser Arg Pro Ser Phe Asp Ala Ile Leu Tyr Tyr Tyr Gln Ser Gly Gly 145 150 155 160

Arg Leu Arg Arg Pro Val Asn Val Pro Leu Asp Val Phe Ser Glu Glu _ 165 170 175

Ile Lys Phe Tyr Glu Leu Gly Asp Gln Ala Ile Asn Lys Phe Arg Glu 180 185 190

Asp Glu Gly Phe Ile Lys Glu Glu Glu Arg Pro Leu Pro Asp Asn Glu 195 200 205

Lys Gln Arg Lys Val Trp Leu Leu Phe Glu Tyr Pro Glu Ser Ser Gln 210 215 220

Ala Ala Arg Val Val Ala Ile Ile Ser Val Phe Val Ile Leu Leu Ser 225 230 235 240

Ile Val Ile Phe Cys Leu Glu Thr Leu Pro Glu Phe Lys His Tyr Lys 245 250 255

Val Phe Asn Thr Thr Asn Gly Thr Lys Ile Glu Glu Asp Glu Val
260 265 270

Pro Asp Ile Thr Asp Pro Phe Phe Leu Ile Glu Thr Leu Cys Ile Ile 275 280 285

Trp Phe Thr Phe Glu Leu Thr Val Arg Phe Leu Ala Cys Pro Asn Lys 290 295 300

Leu Asn Phe Cys Arg Asp Val Met Asn Val Ile Asp Ile Ile Ala Ile 305 310 315 320

Ile Pro Tyr Phe Ile Thr Leu Ala Thr Val Val Ala Glu Glu Asp 325 330 335

Thr Leu Asn Leu Pro Lys Ala Pro Val Ser Pro Gln Asp Lys Ser Ser 340 345 350

Asn Gln Ala Met Ser Leu Ala Ile Leu Arg Val Ile Arg Leu Val Arg 355 360 365

Val Phe Arg Ile Phe Lys Leu Ser Arg His Ser Lys Gly Leu Gln Ile 370 375 380

Leu Gly Arg Thr Leu Lys Ala Ser Met Arg Glu Leu Gly Leu Leu Ile 385 390 395 400

Phe Phe Leu Phe Ile Gly Val Val Leu Phe Ser Ser Ala Val Tyr Phe 405 410 415

Ala Glu Ala Gly Ser Glu Asn Ser Phe Phe Lys Ser Ile Pro Asp Ala 420 425 430

Phe Trp Trp Ala Val Val Thr Met Thr Thr Val Gly Tyr Gly Asp Met 435 440 445

Thr Pro Val Gly Phe Trp Gly Lys Ile Val Gly Ser Leu Cys Val Ile 450 455 460

Ala Gly Val Leu Thr Ile Ala Leu Pro Val Pro Val Ile Val Ser Asn 465 470 475 480

Phe Asn Tyr Phe Tyr His Arg Glu Ala Asp Arg Glu Glu Met Gln Ser 485 490 495

Gln Asn Phe Asn His Val Thr Ser Cys Ser Tyr Leu Pro Gly Ala Leu 500 505 510

Gly Gln His Leu Lys Lys Ser Ser Leu Ser Glu Ser Ser Ser Asp Ile 515 520 525

Met Asp Leu Asp Asp Gly Ile Asp Ala Thr Thr Pro Gly Leu Thr Asp 530 540

His Thr Gly Arg His Met Val Pro Phe Leu Arg Thr Gln Gln Ser Phe 545 550 555 560

Glu Lys Gln Gln Leu Gln Leu Gln Leu Gln Leu Gln Gln Gln Ser Gln 565 570 575

Ser Pro His Gly Gln Gln Met Thr Gln Gln Gln Leu Gly Gln Asn 580 585 590

Gly Leu Arg Ser Thr Asn Ser Leu Gln Leu Arg His Asn Asn Ala Met 595 600 605

Ala Val Ser Ile Glu Thr Asp Val

<210> 5

<211> 495

<212> PRT

<213> Homo sapiens

<400> 5

Met Thr Val Met Ser Gly Glu Asn Val Asp Glu Ala Ser Ala Ala Pro 1 5 10 15 Gly His Pro Gln Asp Gly Ser Tyr Pro Arg Gln Ala Asp His Asp Asp 20 25 30

His Glu Cys Cys Glu Arg Val Val Ile Asn Ile Ser Gly Leu Arg Phe
35 40 45

Glu Thr Gln Leu Lys Thr Leu Ala Gln Phe Pro Asn Thr Leu Leu Gly
50 55 60

Asn Pro Lys Lys Arg Met Arg Tyr Phe Asp Pro Leu Arg Asn Glu Tyr 65 70 75 80

Phe Phe Asp Arg Asn Arg Pro Ser Phe Asp Ala Ile Leu Tyr Tyr Tyr 85 90 95

Gln Ser Gly Gly Arg Leu Arg Arg Pro Val Asn Val Pro Leu Asp Met
100 105 110

Phe Ser Glu Glu Ile Lys Phe Tyr Glu Leu Gly Glu Glu Ala Met Glu
115 120 125

Lys Phe Arg Glu Asp Glu Gly Phe Ile Lys Glu Glu Glu Arg Pro Leu 130 135 140

Pro Glu Lys Glu Tyr Gln Arg Gln Val Trp Leu Leu Phe Glu Tyr Pro 145 150 155 160

Glu Ser Ser Gly Pro Ala Arg Val Ile Ala Ile Val Ser Val Met Val 165 170 175

Ile Leu Ile Ser Ile Val Ile Phe Cys Leu Glu Thr Leu Pro Glu Leu , 180 185 190

Lys Asp Asp Lys Asp Phe Thr Gly Thr Val His Arg Ile Asp Asn Thr
195 200 205

Thr Val Ile Tyr Asn Ser Asn Ile Phe Thr Asp Pro Phe Phe Ile Val 210 215 220

Glu Thr Leu Cys Ile Ile Trp Phe Ser Phe Glu Leu Val Val Arg Phe 225 230 235 240

Phe Ala Cys Pro Ser Lys Thr Asp Phe Phe Lys Asn Ile Met Asn Phe 245 250 255

Ile Asp Ile Val Ala Ile Ile Pro Tyr Phe Ile Thr Leu Gly Thr Glu 260 265 270

Ile Ala Glu Gln Glu Gly Asn Gln Lys Gly Glu Gln Ala Thr Ser Leu 275 280 285

Ala Ile Leu Arg Val Ile Arg Leu Val Arg Val Phe Arg Ile Phe Lys 290 295 300

Leu Ser Arg His Ser Lys Gly Leu Gln Ile Leu Gly Gln Thr Leu Lys 305 310 315 320

Ala Ser Met Arg Glu Leu Gly Leu Leu Ile Phe Phe Leu Phe Ile Gly 325 330 335

Val Ile Leu Phe Ser Ser Ala Val Tyr Phe Ala Glu Ala Glu Glu Ala 340 345 350

Glu Ser His Phe Ser Ser Ile Pro Asp Ala Phe Trp Trp Ala Val Val 355 360 365

Ser Met Thr Thr Val Gly Tyr Gly Asp Met Tyr Pro Val Thr Ile Gly 370 375 380

Gly Lys Ile Val Gly Ser Leu Cys Ala Ile Ala Gly Val Leu Thr Ile 385 390 395 400

Ala Leu Pro Val Pro Val Ile Val Ser Asn Phe Asn Tyr Phe Tyr His
405 410 415

Arg Glu Thr Glu Gly Glu Glu Gln Ala Gln Leu Leu His Val Ser Ser 420 425 430

Pro Asn Leu Ala Ser Asp Ser Asp Leu Ser Arg Arg Ser Ser Ser Thr
435
440
445

Met Ser Lys Tyr Glu Tyr Met Glu Ile Glu Glu Asp Met Asn Asn Ser 450 455 460

Ile Ala His Tyr Arg Gln Val Asn Ile Arg Thr Ala Asn Cys Thr Thr 465 470 475 480

Ala Asn Gln Asn Cys Val Asn Lys Ser Lys Leu Leu Thr Asp Val
485 490 495

<210> 6

<211> 858

<212> PRT

<213> Homo sapiens

<400> 6

Met Pro Ala Gly Met Thr Lys His Gly Ser Arg Ser Thr Ser Ser Leu

1 5 10 15

Pro Pro Glu Pro Met Glu Ile Val Arg Ser Lys Ala Cys Ser Arg Arg 20 25 30

Val Arg Leu Asn Val Gly Gly Leu Ala His Glu Val Leu Trp Arg Thr
. 35 40 45

Leu Asp Arg Leu Pro Arg Thr Arg Leu Gly Lys Leu Arg Asp Cys Asn
50 60

Thr His Asp Ser Leu Leu Glu Val Cys Asp Asp Tyr Ser Leu Asp Asp 65 70 75 80

Asn Glu Tyr Phe Phe Asp Arg His Pro Gly Ala Phe Thr Ser Ile Leu 85 90 95

Asn Phe Tyr Arg Thr Gly Arg Leu His Met Met Glu Glu Met Cys Ala 100 105 110

Leu Ser Phe Ser Gln Glu Leu Asp Tyr Trp Gly Ile Asp Glu Ile Tyr
115 120 125

Leu Glu Ser Cys Cys Gln Ala Arg Tyr His Gln Lys Lys Glu Gln Met
130 135 140

Asn Glu Glu Leu Lys Arg Glu Ala Glu Thr Leu Arg Glu Arg Glu Gly

Glu Glu Phe Asp Asn Thr Cys Cys Ala Glu Lys Arg Lys Lys Leu Trp 165 170 175

Asp Leu Leu Glu Lys Pro Asn Ser Ser Val Ala Ala Lys Ile Leu Ala 180 185 . 190

Ile Ile Ser Ile Met Phe Ile Val Leu Ser Thr Ile Ala Leu Ser Leu
195 200 205

Asn Thr Leu Pro Glu Leu Gln Ser Leu Asp Glu Phe Gly Gln Ser Thr 210 215 220

Asp Asn Pro Gln Leu Ala His Val Glu Ala Val Cys Ile Ala Trp Phe 225 230 235 240

Thr Met Glu Tyr Leu Leu Arg Phe Leu Ser Ser Pro Lys Lys Trp Lys 245 250 255

Phe Phe Lys Gly Pro Leu Asn Ala Ile Asp Leu Leu Ala Ile Leu Pro 260 265 270

Tyr Tyr Val Thr Ile Phe Leu Thr Glu Ser Asn Lys Ser Val Leu Gln 275 280 285

Phe Gln Asn Val Arg Arg Val Val Gln Ile Phe Arg Ile Met Arg Ile , 290 295 300

Leu Arg Ile Leu Lys Leu Ala Arg His Ser Thr Gly Leu Gln Ser Leu 305 310 315 320

Gly Phe Thr Leu Arg Arg Ser Tyr Asn Glu Leu Gly Leu Leu Ile Leu 325 330 335

Phe Leu Ala Met Gly Ile Met Ile Phe Ser Ser Leu Val Phe Phe Ala 340 345 350

Glu Lys Asp Glu Asp Asp Thr Lys Phe Lys Ser Ile Pro Ala Ser Phe 355 360 365

Trp Trp Ala Thr Ile Thr Met Thr Thr Val Gly Tyr Gly Asp Ile Tyr 370 375 380

Pro Lys Thr Leu Leu Gly Lys Ile Val Gly Gly Leu Cys Cys Ile Ala 385 390 395 400

Gly Val Leu Val Ile Ala Leu Pro Ile Pro Ile Ile Val Asn Asn Phe 405 410 415

Ser Glu Phe Tyr Lys Glu Gln Lys Arg Gln Glu Lys Ala Ile Lys Arg 420 425 430

Arg Glu Ala Leu Glu Arg Ala Lys Arg Asn Gly Ser Ile Val Ser Met 435 440 445

Asn Met Lys Asp Ala Phe Ala Arg Ser Ile Glu Met Met Asp Ile Val 450 455 460

Val Glu Lys Asn Gly Glu Asn Met Gly Lys Lys Asp Lys Val Gln Asp 465 470 475 480

Asn His Leu Ser Pro Asn Lys Trp Lys Trp Thr Lys Arg Thr Leu Ser 485 490 495

Glu Thr Ser Ser Ser Lys Ser Phe Glu Thr Lys Glu Gln Gly Ser Pro
500 505 510

Glu Lys Ala Arg Ser Ser Ser Pro Gln His Leu Asn Val Gln Gln
515 520 525

Leu Glu Asp Met Tyr Asn Lys Met Ala Lys Thr Gln Ser Gln Pro Ile 530 535 540

Leu Asn Thr Lys Glu Ser Ala Ala Gln Ser Lys Pro Lys Glu Glu Leu 545 550 555 560

Glu Met Glu Ser Ile Pro Ser Pro Val Ala Pro Leu Pro Thr Arg Thr . 565 570 575

Glu Gly Val Ile Asp Met Arg Ser Met Ser Ser Ile Asp Ser Phe Ile 580 585 590

Ser Cys Ala Thr Asp Phe Pro Glu Ala Thr Arg Phe Ser His Ser Pro 595 600 605

Leu Thr Ser Leu Pro Ser Lys Thr Gly Gly Ser Thr Ala Pro Glu Val , 610 615 620

Gly Trp Arg Gly Ala Leu Gly Ala Ser Gly Gly Arg Phe Val Glu Ala 625 630 635 640

Asn Pro Ser Pro Asp Ala Ser Gln His Ser Ser Phe Phe Ile Glu Ser 645 650 655

Pro Lys Ser Ser Met Lys Thr Asn Asn Pro Leu Lys Leu Arg Ala Leu 660 665 670

Lys Val Asn Phe Met Glu Gly Asp Pro Ser Pro Leu Leu Pro Val Leu 675 680 685

Gly Met Tyr His Asp Pro Leu Arg Asn Arg Gly Ser Ala Ala Ala 690 695 700

Val Ala Gly Leu Glu Cys Ala Thr Leu Leu Asp Lys Ala Val Leu Ser 705 710 715 720

Pro Glu Ser Ser Ile Tyr Thr Thr Ala Ser Ala Lys Thr Pro Pro Arg
725 730 735

Ser Pro Glu Lys His Thr Ala Ile Ala Phe Asn Phe Glu Ala Gly Val 740 745 750

His Gln Tyr Ile Asp Ala Asp Thr Asp Asp Glu Gly Gln Leu Leu Tyr 755 760 765

Ser Val Asp Ser Ser Pro Pro Lys Ser Leu Pro Gly Ser Thr Ser Pro 770 780

Lys Phe Ser Thr Gly Thr Arg Ser Glu Lys Asn His Phe Glu Ser Ser 785 790 795 800

Pro Leu Pro Thr Ser Pro Lys Phe Leu Arg Gln Asn Cys Ile Tyr Ser 805 810 815

Thr Glu Ala Leu Thr Gly Lys Gly Pro Ser Gly Gln Glu Lys Cys Lys 820 825 830

Leu Glu Asn His Ile Ser Pro Asp Val Arg Val Leu Pro Gly Gly Gly 835 840 845

Ala His Gly Ser Thr Arg Asp Gln Ser Ile 850 855

<210> 7

<211> 597

<212> PRT

<213> Paramecium tetraurelia

<400> 7

Met Gly Pro Lys Ile Arg Ala Ile Ser Phe Gln Ser Asn Lys Pro Met
1 5 10 15

Met Asn Leu Lys Glu Asp Ser Pro Val Phe Ile Asp Ser His Thr Asp 20 25 30

His Val Gly Phe Ser Asn Lys Ile Trp Arg Thr Lys Ala Leu Glu Ile 35 40 45

Leu Met Ile Thr Leu Arg Phe Ile Ser Phe Ile Thr Lys Ser Asn Phe 50 55 60

Ala Thr Ser Phe Lys Leu Ile Asn Lys Asn Val Phe Glu Ile Ile Gly 65 70 75 80

Asp Val Ser Ala Asp Phe Thr Tyr Tyr Leu Leu Lys Asn Phe Phe Lys 85 90 95

Tyr Glu Lys Pro Thr Gly Phe Gln Lys Gly Gln His Phe Leu Asn Gln
100 105 110

Thr Leu Leu Ile Pro Leu Arg Lys Thr Lys Leu Leu Lys Ile Tyr Cys
115 120 125

Gly Asn Gln Lys Leu Ile Met Arg Pro Glu Ser Leu Ala Ser Ile Trp 130 135 140

Trp Asn Ile Tyr Ile Leu Thr Ile Leu Asn Ile Asn Val Leu Tyr Val 145 150 155 160

Ser Ile Lys Ile Ala Phe Lys Phe Asp Glu Gln Ser Gln Asp Asp Phe 165 170 175

Tyr Gln Ala Arg Gln Ile Ile Phe Asp Val Leu Pro Ser Tyr Ser Phe 180 185 190

- Met Leu Glu Ile Leu Leu Lys Phe Asn Thr Cys Tyr Tyr Tyr Lys Gly
 195 200 205
- Ala Val Ile Glu Asn Arg Tyr Gln Ile Ala Lys Asn Tyr Leu Arg Ser 210 215 220
- Ser Phe Phe Phe Asp Ile Phe Val Val Ile Pro Tyr Phe Ile Ser Leu 225 230 235 240
- Arg Phe Asp Leu Gln Tyr Leu Asp Leu Val Ile Ile Leu Lys Val Phe 245 250 255
- Gln Ile Thr Lys Phe Ser Arg Asn Leu Phe Asp Arg Leu Glu Leu Thr 260 265 270
- Ala Ile Gln Ile Val Ile Val Asp Leu Val Lys Leu Gly Tyr Thr Ile 275 280 285
- Leu Ala Ala Ala His Phe Ser Ala Cys Ile Trp Phe Leu Val Gly Ser 290 295 300
- Thr Gly Asn Pro Asn Asp Thr Ser Trp Ile Lys Ala Gln Asn Ile Glu 305 310 315 320
- Asn Glu Gln Trp Phe Asn Gln Tyr Leu His Ser Leu Tyr Trp Ser Ile 325 330 335
- Ile Thr Met Thr Thr Ile Gly Tyr Gly Asp Ile Thr Pro Gln Asn Leu 340 345 350
- Arg Glu Arg Val Phe Ala Val Gly Met Ala Leu Ser Ala Val Gly Val , 355 360 365
- Phe Gly Tyr Ser Ile Gly Asn Ile Asn Ser Ile Tyr Ala Glu Trp Ser 370 375 380
- Arg Gln Ser Phe Gln Ile Arg Thr Asp Met Asn Asn Leu Lys Lys Phe 385 390 395 400
- Ile Arg Ile Lys Gly Ile Asn Lys His Leu Ala Glu Lys Ile Arg Lys
 405 410 415
- Tyr Phe Glu Tyr Val Trp Ser Asp Gln Met Glu Asp Asn Asp Arg Glu
 420 425 430
- Val Tyr Lys Phe Ser Glu Met Ile Pro Lys Gln Leu Ala Glu Glu Met 435 440 445
- Lys Ile Asp Thr Asn Met Lys Leu Ile Gln Lys Asn Ser Phe Leu Val 450 455 460
- Asn Asn Phe Ser Glu Gln Phe Leu Ile Ser Leu Ser Lys Val Leu Ile 465 470 475 480
- Glu Glu Lys Tyr Val Pro Glu Ser Thr Ile Tyr Leu Val Lys Leu Ile 485 490 495
- Asn Ile Leu Gln Gln Asn Asp Pro Ser Asn Tyr Leu Tyr Ile Leu Ser 500 505 510

Asn Gly Ser Leu Ser Phe Tyr Ile Thr Leu Asn Asn Lys Gln Gln Thr 515 520 525

Ile Lys Val Leu Glu Thr Ile Lys Asn Glu Gly Gln Ala Phe Gly Val 530 540

Leu Glu Phe Phe Gln Ser Gln Ala Tyr Gln Val Ser Cys Lys Ser Asn 545 550 555 560

Gln Phe Ser Tyr Val Leu Lys Ile Asp Lys Ser Gln Phe Met Glu Ile 565 570 575

Ile Ser Gln His Lys Asn Asp Tyr Val Thr Gln Ile Ile Tyr Leu Ile 580 585 590

Leu Val Gln Ile Leu 595

<210> 8

<211> 556

<212> PRT

<213> Caenorhabditis elegans

<400> 8

Asp Ala Cys Ser Phe Asn Arg Phe Asp Ser Asn Arg Ser Ser Ala Arg

1 5 10 15

Arg Phe Ser Arg Arg Gly Ser Asp Tyr Phe Gly Asp Lys Gly Ile Ser 20 25 30

Met Asp Glu Arg Ile Val Leu Asn Val Gly Gly Val Arg His Glu Thr 35 40 45

Tyr Gln Ala Thr Leu Lys Lys Ile Pro Ala Thr Arg Leu Ser Arg Leu 50 60

Thr Pro Ser Leu Ala Asn Phe Asp Pro Leu Leu Asn Glu Tyr Phe Phe 65 70 75 80

Asp Arg His Pro Ala Val Phe Ala Met Ile Leu Asn Tyr Tyr Arg Thr
85 90 95

Gly Lys Leu His Tyr Pro Thr Asp Val Cys Gly Pro Leu Phe Glu Glu 100 105 110

Glu Leu Gln Tyr Trp Gly Leu Asp Ala Ser Asp Thr Glu Pro Cys Cys 115 120 125

Trp Met Gln Leu Leu His Ala Lys Asp Thr Gln Glu Thr Leu Ala Val 130 135 140

Leu Asp Arg Met Asp Ala Asp His Glu Asp Asp Pro Gln Leu Arg Glu 145 150 155 160

Gln Asp Thr Met Lys Lys Phe Gly Trp Glu Glu Asp Tyr Phe Gln Gly
165 170 175

Lys Arg Thr Arg Trp Met Lys Leu Lys Pro Gln Met Trp Ser Leu Phe 180 185 190

- Asp Glu Pro Tyr Ser Ser Gln Ala Ala Lys Leu Ile Ala Gly Ile Ser 195 200 205
- Val Leu Phe Ile Phe Ile Ser Ile Phe Ser Phe Cys Leu Lys Thr His 210 220
- Gln Ser Phe Arg Leu Pro Val Leu Ile Gly Gln Asn Ile Thr Met Pro 225 230 235 240
- Gly Gly Val Val Gln Pro Ser Ile Glu Arg Val Ser Thr Glu Pro Leu 245 250 255
- Pro Ile Phe Gly Gln Ile Glu Met Leu Cys Asn Ile Trp Phe Thr Leu 260 265 270
- Glu Leu Ile Ile Arg Phe Val Phe Cys Pro Ser Lys Ile Arg Phe Phe 275 280 285
- Lys Ser Pro Leu Asn Met Ile Asp Leu Val Ala Thr Leu Ser Phe Tyr 290 295 300
- Ala Asp Ala Met Met Val Arg Val Val Glu Asp Glu Pro Lys Asp Val 305 310 315 320
- Val Glu Phe Leu Ser Met Ile Arg Ile Phe Arg Leu Phe Lys Leu Thr 325 330 335
- Gln His His Gln Gly Leu Gln Ile Leu Ile His Thr Phe Arg Ala Ser
- Ala Lys Glu Leu Ile Leu Leu Val Phe Phe Leu Ile Leu Gly Ile Val , 355 360 365
- Ile Phe Ala Ala Leu Val Tyr Tyr Ala Glu Lys Met Glu Ala Asn Pro 370 375 380
- Asn Asn Gln Phe Gln Ser Ile Pro Leu Gly Leu Trp Trp Ala Ile Cys 385 390 395 400
- Thr Met Thr Thr Val Gly Tyr Gly Asp Met Thr Pro His Thr Ser Phe 405 · 410 415
- Gly Arg Leu Val Gly Ser Leu Cys Ala Val Met Gly Val Leu Thr Ile 420 425 430
- Ala Leu Pro Val Pro Val Ile Val Ser Asn Phe Ala Met Phe Tyr Ser 435 440 445
- His Asn Gln Ala Arg Asp Lys Leu Pro Lys Arg Arg Arg Val Leu
 450 455 460
- Pro Val Glu Gln Ile Arg Leu Gln Ala Arg Arg His Ala Ala Val Leu 465 470 475 480
- Glu Pro Ser Ala Ser Gln Gly Gly Leu Gly Gly Gln Ala Ile Arg
 485 490 495
- Arg Arg Asn Met Pro Ile Leu Ile Asp Gln Asn Cys Cys Asp Glu Glu
 500 505 510

Asn His Asn His Lys Asp Arg Glu Lys Ser Glu Asn Ser Asp Glu Gly 515 520 525

Thr Asn Ser Ser Ser Thr Thr Gly Val Asp Thr Val Val Lys Leu Gly 530 540

Pro Ser Glu Thr Ala Ile Thr Thr Thr Ile Ile Ser 545 550 555

<210> 9

<211> 1196

<212> PRT

<213> Mus musculus

<400> 9

Met Glu Leu Glu His Pro Lys Ser Pro Pro Tyr Pro Ser Ser Ser 1 5 10 15

Ser Ser Ser Ser Ser Val His Glu Pro Lys Met Asp Ala Leu Ile 20 25 30

Ile Pro Val Thr Met Glu Val Pro Cys Asp Ser Arg Gly Gln Arg Met
35 40 45

Trp Trp Ala Phe Leu Ala Ser Ser Met Val Thr Phe Phe Gly Gly Leu
50 55 60

Phe Ile Ile Leu Leu Trp Arg Thr Leu Lys Tyr Leu Trp Thr Val Cys
65 70 75 80

Cys His Cys Gly Gly Lys Thr Lys Glu Ala Gln Lys Ile Asn Asn Gly 85 90 95

Ser Ser Gln Ala Asp Gly Thr Leu Lys Pro Val Asp Glu Lys Glu Glu
100 105 110

Val Val Ala Ala Glu Val Gly Trp Met Thr Ser Val Lys Asp Trp Ala 115 120 125

Gly Val Met Ile Ser Ala Gln Thr Leu Thr Gly Arg Val Leu Val Val 130 135 140

Leu Val Phe Ala Leu Ser Ile Gly Ala Leu Val Ile Tyr Phe Ile Asp 145 150 155 160

Ser Ser Asn Pro Ile Glu Ser Cys Gln Asn Phe Tyr Lys Asp Phe Thr 165 170 175

Leu Gln Ile Asp Met Ala Phe Asn Val Phe Phe Leu Leu Tyr Phe Gly
180 .185 190

Leu Arg Phe Ile Ala Ala Asn Asp Lys Leu Trp Phe Trp Leu Glu Val 195 200 205

Asn Ser Val Val Asp Phe Phe Thr Val Pro Pro Val Phe Val Ser Val 210 215 220

Tyr Leu Asn Arg Ser Trp Leu Gly Leu Arg Phe Leu Arg Ala Leu Arg 225 230 235 240

- Leu Ile Gln Phe Ser Glu Ile Leu Gln Phe Leu Asn Ile Leu Lys Thr 245 250 255
- Ser Asn Ser Ile Lys Leu Val Asn Leu Leu Ser Ile Phe Ile Ser Thr 260 265 270
- Trp Leu Thr Ala Ala Gly Phe Ile His Leu Val Glu Asn Ser Gly Asp 275 280 285
- Pro Trp Glu Asn Phe Gln Asn Asn Gln Ala Leu Thr Tyr Trp Glu Cys 290 295 300
- Val Tyr Leu Leu Met Val Thr Met Ser Thr Val Gly Tyr Gly Asp Val 305 310 315 320
- Tyr Ala Lys Thr Thr Leu Gly Arg Leu Phe Met Val Phe Phe Ile Leu 325 330 335
- Gly Gly Leu Ala Met Phe Ala Ser Tyr Val Pro Glu Ile Ile Glu Leu 340 345 350
- Ile Gly Asn Arg Lys Lys Tyr Gly Gly Ser Tyr Ser Ala Val Ser Gly 355 360 365
- Arg Lys His Ile Val Val Cys Gly His Ile Thr Leu Glu Ser Val Ser 370 380
- Asn Phe Leu Lys Asp Phe Leu His Lys Asp Arg Asp Asp Val Asn Val 385 390 395 400
- Glu Ile Val Phe Leu His Asn Ile Ser Pro Asn Leu Glu Leu Glu Ala ,
 405 410 415
- Leu Phe Lys Arg His Phe Thr Gln Val Glu Phe Tyr Gln Gly Ser Val
 420 425 430
- Leu Asn Pro His Asp Leu Ala Arg Val Lys Ile Glu Ser Ala Asp Ala
 435 440 445
- Cys Leu Ile Leu Ala Asn Lys Tyr Cys Ala Asp Pro Asp Ala Glu Asp 450 455 460
- Ala Ser Asn Ile Met Arg Val Ile Ser Ile Lys Asn Tyr His Pro Lys 465 470 475 480
- Ile Arg Ile Ile Thr Gln Met Leu Gln Tyr His Asn Lys Ala His Leu 485 490 495
- Leu Asn Ile Pro Ser Trp Asn Trp Lys Glu Gly Asp Asp Ala Ile Cys
 500 505 510
- Leu Ala Glu Leu Lys Leu Gly Phe Ile Ala Gln Ser Cys Leu Ala Gln 515 520 525
- Gly Leu Ser Thr Met Leu Ala Asn Leu Phe Ser Met Arg Ser Phe Ile 530 535 540
- Lys Ile Glu Glu Asp Thr Trp Gln Lys Tyr Tyr Leu Glu Gly Val Ser 545 550 555 560

Asn Glu Met Tyr Thr Glu Tyr Leu Ser Ser Ala Phe Val Gly Leu Ser Phe Pro Thr Val Cys Glu Leu Cys Phe Val Lys Leu Lys Leu Leu Met 580 Ile Ala Ile Glu Tyr Lys Ser Ala Asn Arg Glu Ser Arg Ile Leu Ile Asn Pro Gly Asn His Leu Lys Ile Gln Glu Gly Thr Leu Gly Phe Phe 610 Ile Ala Ser Asp Ala Lys Glu Val Lys Arg Ala Phe Phe Tyr Cys Lys 630 Ala Cys His Asp Asp Val Thr Asp Pro Lys Arg Ile Lys Lys Cys Gly 645 Cys Arg Arg Leu Ile Tyr Phe Glu Asp Glu Gln Pro Pro Thr Leu Ser 665 Pro Lys Lys Gln Arg Asn Gly Gly Met Arg Asn Ser Pro Asn Thr 675 680 Ser Pro Lys Leu Met Arg His Asp Pro Leu Leu Ile Pro Gly Asn Asp 695 Gln Ile Asp Asn Met Asp Ser Asn Val Lys Lys Tyr Asp Ser Thr Gly 715 710 Met Phe His Trp Cys Ala Pro Lys Glu Ile Glu Lys Val Ile Leu Thr 725 Arg Ser Glu Ala Ala Met Thr Val Leu Ser Gly His Val Val Cys 740 745 Ile Phe Gly Asp Val Ser Ser Ala Leu Ile Gly Leu Arg Asn Leu Val Met Pro Leu Arg Ala Ser Asn Phe His Tyr His Glu Leu Lys His Ile 775 Val Phe Val Gly Ser Ile Glu Tyr Leu Lys Arg Glu Trp Glu Thr Leu . 790 His Asn Phe Pro Lys Val Ser Ile Leu Pro Gly Thr Pro Leu Ser Arg 805 Ala Asp Leu Arg Ala Val Asn Ile Asn Leu Cys Asp Met Cys Val Ile 825 Leu Ser Ala Asn Gln Asn Asn Ile Asp Asp Thr Ser Leu Gln Asp Lys 840 835 Glu Cys Ile Leu Ala Ser Leu Asn Ile Lys Ser Met Gln Phe Asp Asp 855 Ser Ile Gly Val Leu Gln Ala Asn Ser Gln Gly Phe Thr Pro Pro Gly 880

875

- Met Asp Arg Ser Ser Pro Asp Asn Ser Pro Val His Gly Met Leu Arg 885 890 895
- Gln Pro Ser Ile Thr Thr Gly Val Asn Ile Pro Ile Ile Thr Glu Leu 900 905 910
- Val Asn Asp Thr Asn Val Gln Phe Leu Asp Gln Asp Asp Asp Asp Asp 915 920 925
- Pro Asp Thr Glu Leu Tyr Leu Thr Gln Pro Phe Ala Cys Gly Thr Ala 930 935 940
- Phe Ala Val Ser Val Leu Asp Ser Leu Met Ser Ala Thr Tyr Phe Asn 945 950 955 960
- Asp Asn Ile Leu Thr Leu Ile Arg Thr Leu Val Thr Gly Gly Ala Thr 965 970 975
- Pro Glu Leu Glu Ala Leu Ile Ala Glu Glu Asn Ala Leu Arg Gly Gly 980 985 990
- Tyr Ser Thr Pro Gln Thr Leu Ala Asn Arg Asp Arg Cys Arg Val Ala 995 1000 1005
- Gln Leu Ala Leu Leu Asp Gly Pro Phe Ala Asp Leu Gly Asp Gly Gly 1010 1015 1020
- Cys Tyr Gly Asp Leu Phe Cys Lys Ala Leu Lys Thr Tyr Asn Met Leu 1025 1030 1035 1040
- Cys Phe Gly Ile Tyr Arg Leu Arg Asp Ala His Leu Ser Thr Pro Ser $_{_{/}}$ 1045 1050 1055
- Gln Cys Thr Lys Arg Tyr Val Ile Thr Asn Pro Pro Tyr Glu Phe Glu 1060 1065 1070
- Leu Val Pro Thr Asp Leu Ile Phe Cys Leu Met Gln Phe Asp His Asn 1075 1080 1085
- Ala Gly Gln Ser Arg Ala Ser Leu Ser His Ser Ser His Ser Ser Gln 1090 1095 1100
- Ser Ser Ser Lys Lys Ser Ser Ser Val His Ser Ile Pro Ser Thr Ala 1105 1110 1115 1120
- Asn Arg Pro Asn Arg Pro Lys Ser Arg Glu Ser Arg Asp Lys Gln Asn 1125 1130 1135
- Ala Thr Arg Met Thr Arg Met Gly Gln Ala Glu Lys Lys Trp Phe Thr 1140 1145 1150
- Asp Glu Pro Asp Asn Ala Tyr Pro Arg Asn Ile Gln Ile Lys Pro Met 1155 1160 1165
- Ser Thr His Met Ala Asn Gln Ile Asn Gln Tyr Lys Ser Thr Ser Ser 1170 1175 1180
- Leu Ile Pro Pro Ile Arg Glu Val Glu Asp Glu Cys 1185 1190 1195

<210> 10 <211> 731 <212> PRT

<213> Homo sapiens

<400> 10

Met Asp Thr Ser Gly His Phe His Asp Ser Gly Val Gly Asp Leu Asp
1 5 10 15

Glu Asp Pro Lys Cys Pro Cys Pro Ser Ser Gly Asp Glu Gln Gln Gln 20 25 30

Gln Gln Gln Gln Gln Gln Gln Pro Pro Pro Ala Ser Pro
35 40 45

Ala Ala Pro Gln Gln Pro Leu Gly Pro Ser'Leu Gln Pro Gln Pro Pro 50 55 60

Ser Pro His Pro Leu Ser Gln Leu Ala Gln Leu Gln Ser Gln Pro Val 85 90 95

His Pro Gly Leu Leu His Ser Ser Pro Thr Ala Phe Arg Ala Pro Pro 100 105 110

Ser Ser Asn Ser Thr Ala Ile Leu His Pro Ser Ser Arg Gln Gly Ser 115 120 125

Gln Leu Asn Leu Asn Asp His Leu Leu Gly His Ser Pro Ser Ser Thr 130 135 140

Ala Thr Ser Gly Pro Gly Gly Gly Ser Arg His Arg Gln Ala Ser Pro 145 150 155 160

Leu Val His Arg Arg Asp Ser Asn Pro Phe Thr Glu Ile Ala Met Ser 165 170 175

Ser Cys Lys Tyr Ser Gly Gly Val Met Lys Pro Leu Ser Arg Phe Ser 180 185 190

Ala Ser Arg Arg Asn Leu Ile Glu Ala Glu Thr Glu Gly Gln Pro Leu 195 200 205

Gln Leu Phe Ser Pro Ser Asn Pro Pro Glu Ile Val Ile Ser Ser Arg 210 215 220

Glu Asp Asn His Ala His Gln Thr Leu Leu His His Pro Asn Ala Thr 225 230 235 240

His Asn His Gln His Ala Gly Thr Thr Ala Ser Ser Thr Thr Phe Pro 245 250 255

Lys Ala Asn Lys Arg Lys Asn Gln Asn Ile Gly Tyr Lys Leu Gly His 260 265 270

Arg Arg Ala Leu Phe Glu Lys Arg Lys Arg Leu Ser Asp Tyr Ala Leu 275 280 285 Ile Phe Gly Met Phe Gly Ile Val Val Met Val Ile Glu Thr Glu Leu 290 295 300

Ser Trp Gly Leu Tyr Ser Lys Asp Ser Met Phe Ser Leu Ala Leu Lys 305 310 315

Cys Arg Ile Ser Leu Ser Thr Ile Ile Leu Leu Gly Leu Ile Ile Ala 325 330 335

Tyr His Thr Arg Gly Val Gln Leu Phe Val Ile Asp Asn Asp Ala Asp 340 345 350

Asp Trp Arg Ile Ala Met Thr Tyr Glu Arg Ile Leu Tyr Ile Ser Leu 355 360 365

Glu Met Leu Val Tyr Thr Asn His Thr Ile Pro Gly Glu Tyr Lys Phe 370 380

Phe Trp Ala Ala Arg Leu Ala Phe Ser Tyr Thr Pro Ser Arg Ala Glu 385 390 395 400

Ala Asp Val Asp Ile Ile Leu Ser Ile Pro Met Phe Leu Arg Leu Tyr 405 410 415

Leu Ile Ala Arg Val Met Leu Leu His Ser Lys Leu Phe Thr Asp Ala 420 425 430

Ser Ser Arg Ser Ile Gly Ala Leu Asn Lys Ile Asn Phe Asn Thr Arg
435
440
445

Phe Val Met Lys Thr Leu Met Thr Ile Cys Pro Gly Thr Val Leu Leu , 450 455 460

Val Phe Ser Ile Ser Leu Trp Ile Ile Ala Ala Trp Thr Val Arg Val 465 470 475 480

Cys Glu Arg Tyr His Asp Gln Gln Asp Val Thr Ser Asn Phe Leu Gly
485 490 495

Ala Met Trp Leu Ile Ser Ile Thr Phe Leu Ser Ile Gly Tyr Gly Asp 500 505 510

Met Val Pro His Thr Tyr Cys Gly Lys Gly Val Cys Leu Leu Thr Gly 515 520 525 .

Ile Met Gly Ala Gly Cys Thr Ala Leu Val Val Ala Val Val Ala Arg 530 535 540

Lys Leu Glu Leu Thr Lys Ala Glu Lys His Val Asp Asn Phe Met Met 545 550 555 560

Asp Thr Gln Leu Thr Lys Arg Ile Lys Asn Ala Ala Ala Asn Val Leu 565 570 575

Arg Glu Thr Trp Leu Ile Tyr Lys His Thr Lys Leu Leu Lys Lys Ile 580 585 590

Asp His Ala Lys Val Arg Lys His Gln Arg Lys Phe Leu Gln Ala Ile 595 600 605 His Gln Leu Arg Ser Val Lys Met Glu Gln Arg Lys Leu Ser Asp Gln 610 620

Ala Asn Thr Leu Val Asp Leu Ser Lys Met Gln Asn Val Met Tyr Asp 625 630 635 640

Leu Ile Thr Glu Leu Asn Asp Arg Ser Glu Asp Leu Glu Lys Gln Ile 645 650 655

Gly Ser Leu Glu Ser Lys Leu Glu His Leu Thr Ala Ser Phe Asn Ser 660 665 670

Leu Pro Leu Leu Ile Ala Asp Thr Leu Arg Gln Gln Gln Gln Gln Leu 675 680 685

Leu Ser Ala Ile Ile Glu Ala Arg Gly Val Ser Val Ala Val Gly Thr 690 695 700

Thr His Thr Pro Ile Ser Asp Thr Pro Ile Gly Val Ser Ser Thr Ser 705 710 715 720

Phe Pro Thr Pro Tyr Thr Ser Ser Ser Cys
725 730

<210> 11

<211> 857

<212> PRT

<213> Arabidopsis thaliana

<400> 11

Met Arg Gly Gly Ala Leu Leu Cys Gly Gln Val Gln Asp Glu Ile Glu
1 5 10 15

Gln Leu Ser Arg Glu Ser Ser His Phe Ser Leu Ser Thr Gly Ile Leu 20 25 30

Pro Ser Leu Gly Ala Arg Ser Asn Arg Arg Val Lys Leu Arg Arg Phe 35 40 45

Val Val Ser Pro Tyr Asp His Lys Tyr Arg Ile Trp Glu Ala Phe Leu
50 55 60

Val Val Leu Val Val Tyr Thr Ala Trp Val Ser Pro Phe Glu Phe Gly 65 70 75 80

Phe Leu Arg Lys Pro Arg Pro Pro Leu Ser Ile Thr Asp Asn Ile Val 85 90 95

Asn Ala Phe Phe Ala Ile Asp Ile Ile Met Thr Phe Phe Val Gly Tyr 100 105 110

Leu Asp Lys Ser Thr Tyr Leu Ile Val Asp Asp Arg Lys Gln Ile Ala 115 120 125

Phe Lys Tyr Leu Arg Ser Trp Phe Leu Leu Asp Leu Val Ser Thr Ile 130 135 140

Pro Ser Glu Ala Ala Met Arg Ile Ser Ser Gln Ser Tyr Gly Leu Phe 145 150 155 160

- Asn Met Leu Arg Leu Trp Arg Leu Arg Arg Val Gly Ala Leu Phe Ala 165 170 175
- Arg Leu Glu Lys Asp Arg Asn Phe Asn Tyr Phe Trp Val Arg Cys Ala 180 185 190
- Lys Leu Val Cys Val Thr Leu Phe Ala Val His Cys Ala Ala Cys Phe 195 200 205
- Tyr Tyr Leu Ile Ala Ala Arg Asn Ser Asn Pro Ala Lys Thr Trp Ile 210 215 220
- Gly Ala Asn Val Ala Asn Phe Leu Glu Glu Ser Leu Trp Met Arg Tyr 225 230 235 240
- Val Thr Ser Met Tyr Trp Ser Ile Thr Thr Leu Thr Thr Val Gly Tyr
 245 250 255
- Gly Asp Leu His Pro Val Asn Thr Lys Glu Met Ile Phe Asp Ile Phe 260 265 270
- Tyr Met Leu Phe Asn Leu Gly Leu Thr Ala Tyr Leu Ile Gly Asn Met 275 280 285
- Thr Asn Leu Val Val His Gly Thr Ser Arg Thr Arg Asn Phe Arg Asp 290 295 300
- Thr Ile Gln Ala Ala Ser Asn Phe Ala His Arg Asn His Leu Pro Pro 305 310 315 320
- Arg Leu Gln Asp Gln Met Leu Ala His Leu Cys Leu Lys Tyr Arg Thr , 325 330 335
- Asp Ser Glu Gly Leu Gln Gln Glu Thr Leu Asp Ala Leu Pro Lys 340 345 350
- Ala Ile Arg Ser Ser Ile Ser His Phe Leu Phe Tyr Ser Leu Met Asp 355 360 365
- Lys Val Tyr Leu Phe Arg Gly Val Ser Asn Asp Leu Leu Phe Gln Leu 370 375 380
- Val Ser Glu Met Lys Ala Glu Tyr Phe Pro Pro Lys Glu Asp Val Ile 385 390 395 400
- Leu Gln Asn Glu Ala Pro Thr Asp Phe Tyr Ile Leu Val Asn Gly Thr 405 410 415
- Ala Asp Leu Val Asp Val Asp Thr Gly Thr Glu Ser Ile Val Arg Glu
 420 425 430
- Val Lys Ala Gly Asp Ile Ile Gly Glu Ile Gly Val Leu Cys Tyr Arg 435 440 445
- Pro Gln Leu Phe Thr Val Arg Thr Lys Arg Leu Cys Gln Leu Leu Arg 450 455 460
- Met Asn Arg Thr Thr Phe Leu Asn Ile Ile Gln Ala Asn Val Gly Asp 465 470 475 480

- Gly Thr Ile Ile Met Asn Asn Leu Leu Gln His Leu Lys Glu Met Asn 485 490 495
- Asp Pro Val Met Thr Asn Val Leu Leu Glu Ile Glu Asn Met Leu Ala 500 505 510
- Arg Gly Lys Met Asp Leu Pro Leu Asn Leu Cys Phe Ala Ala Ile Arg 515 520 525
- Glu Asp Asp Leu Leu Leu His Gln Leu Leu Lys Arg Gly Leu Asp Pro
- Asn Glu Ser Asp Asn Asn Gly Arg Thr Pro Leu His Ile Ala Ala Ser 545 550 555 560
- Lys Gly Thr Leu Asn Cys Val Leu Leu Leu Glu Tyr His Ala Asp 565 570 575
- Pro Asn Cys Arg Asp Ala Glu Gly Ser Val Pro Leu Trp Glu Ala Met 580 585 590
- Val Glu Gly His Glu Lys Val Val Leu Leu Glu His Gly Ser
 595 600 605
- Thr Ile Asp Ala Gly Asp Val Gly His Phe Ala Cys Thr Ala Ala Glu 610 615 620
- Gln Gly Asn Leu Lys Leu Leu Lys Glu Ile Val Leu His Gly Gly Asp 625 630 635 640
- Val Thr Arg Pro Arg Ala Thr Gly Thr Ser Ala Leu His Thr Ala Val , 645 650 655
- Cys Glu Glu Asn Ile Glu Met Val Lys Tyr Leu Leu Glu Gln Gly Ala 660 665 670
- Asp Val Asn Lys Gln Asp Met His Gly Trp Thr Pro Arg Asp Leu Ala 675 680 685
- Glu Gln Gln Gly His Glu Asp Ile Lys Ala Leu Phe Arg Glu Lys Leu 690 695 700
- His Glu Arg Arg Val His Ile Glu Thr Ser Ser Ser Val Pro Ile Leu 705 710 715 720
- Lys Thr Gly Ile Arg Phe Leu Gly Arg Phe Thr Ser Glu Pro Asn Ile 725 730 735
- Arg Pro Ala Ser Arg Glu Val Ser Phe Arg Ile Arg Glu Thr Arg Ala
 740 745 . 750
- Arg Arg Lys Thr Asn Asn Phe Asp Asn Ser Leu Phe Gly Ile Leu Ala 755 760 765
- Asn Gln Ser Val Pro Lys Asn Gly Leu Ala Thr Val Asp Glu Gly Arg 770 775 780
- Thr Gly Asn Pro Val Arg Val Thr Ile Ser Cys Ala Glu Lys Asp Asp 785 790 795 800

Ile Ala Gly Lys Leu Val Leu Leu Pro Gly Ser Phe Lys Glu Leu Leu 805 810 815

Glu Leu Gly Ser Asn Lys Phe Gly Ile Val Ala Thr Lys Val Met Asn 820 825 830

Lys Asp Asn Asn Ala Glu Ile Asp Asp Val Asp Val Ile Arg Asp Gly 835 840 845

Asp His Leu Ile Phe Ala Thr Asp Ser 850 855

<210> 12

<211> 1159

<212> PRT

<213> Homo sapiens

<400> 12

Met Pro Val Arg Arg Gly His Val Ala Pro Gln Asn Thr Phe Leu Asp
1 5 10 15

Thr Ile Ile Arg Lys Phe Glu Gly Gln Ser Arg Lys Phe Ile Ile Ala 20 25 30

Asn Ala Arg Val Glu Asn Cys Ala Val Ile Tyr Cys Asn Asp Gly Phe 35 40 45

Cys Glu Leu Cys Gly Tyr Ser Arg Ala Glu Val Met Gln Arg Pro Cys
50 60

Thr Cys Asp Phe Leu His Gly Pro Arg Thr Gln Arg Arg Ala Ala 65 70 75 80

Gln Ile Ala Gln Ala Leu Leu Gly Ala Glu Glu Arg Lys Val Glu Ile 85 90 95

Ala Phe Tyr Arg Lys Asp Gly Ser Cys Phe Leu Cys Leu Val Asp Val
100 105 110

Val Pro Val Lys Asn Glu Asp Gly Ala Val Ile Met Phe Ile Leu Asn 115 120 125

Phe Glu Val Val Met Glu Lys Asp Met Val Gly Ser Pro Ala His Asp 130 135 140

Thr Asn His Arg Gly Pro Pro Thr Ser Trp Leu Ala Pro Gly Arg Ala
145 150 155 160

Lys Thr Phe Arg Leu Lys Leu Pro Ala Leu Leu Ala Leu Thr Ala Arg 165 170 175

Glu Ser Ser Val Arg Ser Gly Gly Ala Gly Ala Gly Ala Pro Gly
180 185 190

Ala Val Val Val Asp Val Asp Leu Thr Pro Ala Ala Pro Ser Ser Glu
195 200 205

Ser Leu Ala Leu Asp Glu Val Thr Ala Met Asp Asn His Val Ala Gly 210 215 220

225	GIY	PLO	Ala	Giu	230	n. g	213	AIU	Deu	235	Gly	110	O. J		240
Pro	Arg	Ser	Ala	Pro 245	Gly	Gln	Leu	Pro	Ser 250	Pro	Arg	Ala	His	Ser 255	Leu
Asn	Pro	Asp	Ala 260	Ser	Gly	Ser	Ser	Cys 265	Ser	Leu	Ala	Arg	Thr 270	Arg	Ser
Arg	Glu	Ser 275	Cys	Ala	Ser	Val	Arg 280	Arg	Ala	Ser	Ser	Ala 285	Asp	Asp	Ile
Glu	Ala 290	Met	Arg	Ala	Gly	Val 295	Leu	Pro	Pro	Pro	Pro 300	Arg	His	Ala	Ser
Thr 305	Gly	Ala	Met	His	Pro 310	Leu	Arg	Ser	Gly	Leu 315	Leu	Asn	Ser	Thr	Ser 320
Asp	Ser	Asp	Leu	Val 325	Arg	Tyr	Arg	Thr	Ile 330	Ser	Lys	Ile	Pro	Gln 335	Ile
Thr	Leu	Asn	Phe 340	Val	Asp	Leu	Lys	Gly 345	Asp	Pro	Phe	Leu	Ala 350	Ser	Pro
Thr	Ser	Asp 355	Arg	Glu	Ile	Ile	Ala 360	Pro	Lys	Ile	Lys	Glu 365	Arg	Thr	His
Asn	Val 370	Thr	Glu	Lys	Val	Thr 375	Gln	Val	Leu	Ser	Leu 380	Gly	Ala	Asp	Val
Leu 385	Pro	Glu	Tyr	Lys	Leu 390	Gln	Ala	Pro	Arg	Ile 395	His	Arg	Trp	Thr	Ile 400
Leu	His	Tyr	Ser	Pro 405	Phe	Lys	Ala	Val	Trp 410	Asp	Trp	Leu	Ile	Leu 415	Leu
Leu	Val	Ile	Tyr 420	Thr	Ala	Val	Phe	Thr 425	Pro	Tyr	Ser	Ala	Ala 430	Phe	Leu
Leu	Lys	Glu 435	Thr	Glu	Glu	Gly	Pro 440	Pro	Ala	Thr	Glu	Cys 445	Gly	Tyr	Ala
Cys	Gln 450	Pro	Leu	Ala	Val	Val 455	Asp	Leu	Ile	Val	Asp 460	Ile	Met	Phe	Ile
Val 465	Asp	Ile	Leu	Ile	Asn 470	Phe	Arg	Thr	Thr	Tyr 475	Val	Asn	Ala	Asn	Glu 480
Glu	Val	Val	Ser	His 485	Pro	Gly	Arg	Ile	Ala 490	Val	His	туr	Phe	Lys 495	Gly
Trp	Phe	Leu	Ile 500	Asp	Met	Val	Ala	Ala 505	Ile	Pro	Phe	Asp	Leu 510	Leu	Ile
Phe	Gly	Ser 515	Gly	Ser	Glu	Glu	Leu 520	Ile	Gly	Leu	Leu	Lys 525	Thr	Ala	Arg
Leu	Leu 530	Arg	Leu	Val	Arg	Val 535	Ala	Arg	Lys	Leu	Asp 540	Arg	Tyr	Ser	Glu

- Tyr Gly Ala Ala Val Leu Phe Leu Leu Met Cys Thr Phe Ala Leu Ile 545 550 555 560
- Ala His Trp Leu Ala Cys Ile Trp Tyr Ala Ile Gly Asn Met Glu Gln 565 570 575
- Pro His Met Asp Ser Arg Ile Gly Trp Leu His Asn Leu Gly Asp Gln
 580 585 590
- Ile Gly Lys Pro Tyr Asn Ser Ser Gly Leu Gly Gly Pro Ser Ile Lys
 595 600 605
- Asp Lys Tyr Val Thr Ala Leu Tyr Phe Thr Phe Ser Ser Leu Thr Ser 610 620
- Val Gly Phe Gly Asn Val Ser Pro Asn Thr Asn Ser Glu Lys Ile Phe 625 630 635 640
- Ser Ile Cys Val Met Leu Ile Gly Ser Leu Met Tyr Ala Ser Ile Phe 645 650 655
- Gly Asn Val Ser Ala Ile Ile Gln Arg Leu Tyr Ser Gly Thr Ala Arg
 660 665 670
- Tyr His Thr Gln Met Leu Arg Val Arg Glu Phe Ile Arg Phe His Gln 675 680 685
- Ile Pro Asn Pro Leu Arg Gln Arg Leu Glu Glu Tyr Phe Gln His Ala 690 695 700
- Trp Ser Tyr Thr Asn Gly Ile Asp Met Asn Ala Val Leu Lys Gly Phe 705 710 715 720
- Pro Glu Cys Leu Gln Ala Asp Ile Cys Leu His Leu Asn Arg Ser Leu 725 730 735
- Leu Gln His Cys Lys Pro Phe Arg Gly Ala Thr Lys Gly Cys Leu Arg
 740 745 750
- Ala Leu Ala Met Lys Phe Lys Thr Thr His Ala Pro Pro Gly Asp Thr 755 760 765
- Leu Val His Ala Gly Asp Leu Leu Thr Ala Leu Tyr Phe Ile Ser Arg 770 775 780
- Gly Ser Ile Glu Ile Leu Arg Gly Asp Val Val Val Ala Ile Leu Gly
 785 790 795 800
- Lys Asn Asp Ile Phe Gly Glu Pro Leu Asn Leu Tyr Ala Arg Pro Gly 805 810 815
- Lys Ser Asn Gly Asp Val Arg Ala Leu Thr Tyr Cys Asp Leu His Lys 820 825 830
- Ile His Arg Asp Asp Leu Leu Glu Val Leu Asp Met Tyr Pro Glu Phe 835 840 845
- Ser Asp His Phe Trp Ser Ser Leu Glu Ile Thr Phe Asn Leu Arg Asp 850 860

- Thr Asn Met Ile Pro Gly Ser Pro Gly Ser Thr Glu Leu Glu Gly Gly 865 870 875 880
- Phe Ser Arg Gln Arg Lys Arg Lys Leu Ser Phe Arg Arg Arg Thr Asp 885 890 895
- Lys Asp Thr Glu Gln Pro Gly Glu Val Ser Ala Leu Gly Pro Gly Arg
 900 905 910
- Ala Gly Ala Gly Pro Ser Ser Arg Gly Arg Pro Gly Gly Pro Trp Gly
 915 920 925
- Glu Ser Pro Ser Ser Gly Pro Ser Ser Pro Glu Ser Ser Glu Asp Glu 930 935 940
- Gly Pro Gly Arg Ser Ser Ser Pro Leu Arg Leu Val Pro Phe Ser Ser 945 955 960
- Pro Arg Pro Pro Gly Glu Pro Pro Gly Glu Pro Leu Met Glu Asp 965 970 975
- Cys Glu Lys Ser Ser Asp Thr Cys Asn Pro Leu Ser Gly Ala Phe Ser 980 985 990
- Gly Val Ser Asn Ile Phe Ser Phe Trp Gly Asp Ser Arg Gly Arg Gln 995 1000 1005
- Tyr Gln Glu Leu Pro Arg Cys Pro Ala Pro Thr Pro Ser Leu Leu Asn 1010 1015 1020
- Ile Pro Leu Ser Ser Pro Gly Arg Arg Pro Arg Gly Asp Val Glu Ser 1025 1030 1035 1040
- Arg Leu Asp Ala Leu Gln Arg Gln Leu Asn Arg Leu Glu Thr Arg Leu 1045 1050 1055
- Ser Ala Asp Met Ala Thr Val Leu Gln Leu Gln Arg Gln Met Thr 1060 1065 1070
- Leu Val Pro Pro Ala Tyr Ser Ala Val Thr Thr Pro Gly Pro Gly Pro 1075 1080 1085
- Thr Ser Thr Ser Pro Leu Leu Pro Val Ser Pro Leu Pro Thr Leu Thr 1090 1095 1100
- Leu Asp Ser Leu Ser Gln Val Ser Gln Phe Met Ala Cys Glu Glu Leu 1105 1110 1115 1120
- Pro Pro Gly Ala Pro Glu Leu Pro Gln Glu Gly Pro Thr Arg Arg Leu.
 1125 1130 1135
- Ser Leu Pro Gly Gln Leu Gly Ala Leu Thr Ser Gln Pro Leu His Arg 1140 1145 1150
- His Gly Ser Asp Pro Gly Ser 1155

<210> 13

<211> 391

<212> PRT

<213> Rattus norvegicus

<400> 13

Met Gly Ala Ser Glu Arg Ser Val Phe Arg Val Leu Ile Arg Ala Leu 1 5 10 15

Thr Glu Arg Met Phe Lys His Leu Arg Arg Trp Phe Ile Thr His Ile 20 25 30

Phe Gly Arg Ser Arg Gln Arg Ala Arg Leu Val Ser Lys Glu Gly Arg
35 40 45

Cys Asn Ile Glu Phe Gly Asn Val Asp Ala Gln Ser Arg Phe Ile Phe 50 55 60

Phe Val Asp Ile Trp Thr Thr Val Leu Asp Leu Lys Trp Arg Tyr Lys
65 70 75 80

Met Thr Val Phe Ile Thr Ala Phe Leu Gly Ser Trp Phe Leu Phe Gly 85 90 95

Leu Leu Trp Tyr Val Val Ala Tyr Val His Lys Asp Leu Pro Glu Phe 100 105 110

Tyr Pro Pro Asp Asn Arg Thr Pro Cys Val Glu Asn Ile Asn Gly Met
115 120 . 125

Thr Ser Ala Phe Leu Phe Ser Leu Glu Thr Gln Val Thr Ile Gly Tyr 130 135 140

Gly Phe Arg Phe Val Thr Glu Gln Cys Ala Thr Ala Ile Phe Leu Leu 145 150 155 160

Ile Phe Gln Ser Ile Leu Gly Val Ile Ile Asn Ser Phe Met Cys Gly
165 170 175

Ala Ile Leu Ala Lys Ile Ser Arg Pro Lys Lys Arg Ala Lys Thr Ile 180 185 190

Thr Phe Ser Lys Asn Ala Val Ile Ser Lys Arg Gly Gly Lys Leu Cys 195 200 205

Leu Leu Ile Arg Val Ala Asn Leu Arg Lys Ser Leu Leu Ile Gly Ser 210 215 220

His Ile Tyr Gly Lys Leu Leu Lys Thr Thr Ile Thr Pro Glu Gly Glu 225 230 235 240

Thr Ile Ile Leu Asp Gln Thr Asn Ile Asn Phe Val Val Asp Ala Gly
245 250 255

Asn Glu Asn Leu Phe Phe Ile Ser Pro Leu Thr Ile Tyr His Ile Ile

Asp His Asn Ser Pro Phe Phe His Met Ala Ala Glu Thr Leu Ser Gln 275 280 285 Gln Asp Phe Glu Leu Val Val Phe Leu Asp Gly Thr Val Glu Ser Thr 290 295 300

Ser Ala Thr Cys Gln Val Arg Thr Ser Tyr Val Pro Glu Glu Val Leu 305 310 315 320

Trp Gly Tyr Arg Phe Val Pro Ile Val Ser Lys Thr Lys Glu Gly Lys
325
330
335

Tyr Arg Val Asp Phe His Asn Phe Gly Lys Thr Val Glu Val Glu Thr 340 345 350

Pro His Cys Ala Met Cys Leu Tyr Asn Glu Lys Asp Ala Arg Ala Arg 355 360 365

Met Lys Arg Gly Tyr Asp Asn Pro Asn Phe Val Leu Ser Glu Val Asp 370 375 380

Glu Thr Asp Asp Thr Gln Met 385 390

<210> 14

<211> 407

<212> PRT

<213> Homo sapiens

<400> 14

Met Asp Gln Asp Val Glu Ser Pro Val Ala Ile His Gln Pro Lys Leu
1 5 10 15

Pro Lys Gln Ala Arg Asp Asp Leu Pro Arg His Ile Ser Arg Asp Arg 20 25 30

Thr Lys Arg Lys Ile Gln Arg Tyr Val Arg Lys Asp Gly Lys Cys Asn 35 40 45

Val His His Gly Asn Val Arg Glu Thr Tyr Arg Tyr Leu Thr Asp Ile 50 55 60

Phe Thr Thr Leu Val Asp Leu Lys Trp Arg Phe Asn Leu Leu Ile Phe 65 70 75 80

Val Met Val Tyr Thr Val Thr Trp Leu Phe Phe Gly Met Ile Trp Trp
85 90 95

Leu Ile Ala Tyr Ile Arg Gly Asp Met Asp His Ile Glu Asp Ser Pro 100 105 110

Trp Thr Pro Cys Val Thr Asn Leu Asn Gly Phe Val Ser Ala Phe Leu 115 120 125

Phe Ser Ile Glu Thr Glu Thr Thr Ile Gly Tyr Gly Tyr Arg Val Ile 130 \$135\$ 140

Thr Asp Lys Cys Pro Glu Gly Ile Ile Leu Leu Leu Ile Gln Ser Val 145 150 155 160

Leu Gly Ser Ile Val Asn Ala Phe Met Val Gly Cys Met Phe Val Lys 165 170 175 Ile Ser Gln Pro Lys Lys Arg Ala Glu Thr Leu Val Phe Ser Thr His 180 185 190

Ala Val Ile Ser Met Arg Asp Gly Lys Leu Cys Leu Met Phe Arg Val 195 200 205

Gly Asp Leu Arg Asn Ser His Ile Val Glu Ala Ser Ile Arg Ala Lys 210 215 220

Leu Ile Lys Ser Lys Gln Thr Ser Glu Gly Glu Phe Ile Pro Leu Asn 225 230 235 240

Gln Thr Asp Ile Asn Val Gly Tyr Tyr Thr Gly Asp Asp Arg Leu Phe 245 250 255

Leu Val Ser Pro Leu Ile Ile Ser His Glu Ile Asn Gln Gln Ser Pro 260 265 270

Phe Trp Glu Ile Ser Lys Ala Gln Leu Pro Lys Glu Glu Leu Glu Ile 275 280 285

Val Val Ile Leu Glu Gly Met Val Glu Ala Thr Gly Met Thr Cys Gln 290 295 300

Ala Arg Ser Ser Tyr Ile Thr Ser Glu Ile Leu Trp Gly Tyr Arg Phe 305 310 315 320

Thr Pro Val Leu Thr Leu Glu Asp Gly Phe Tyr Glu Val Asp Tyr Asn 325 330 335

Ser Phe His Glu Thr Tyr Glu Thr Ser Thr Pro Ser Leu Ser Ala Lys 340 345 350

Glu Leu Ala Glu Leu Ala Ser Arg Ala Glu Leu Pro Leu Ser Trp Ser 355 360 365

Val Ser Ser Lys Leu Asn Gln His Ala Glu Leu Glu Thr Glu Glu Glu 370 380

Glu Lys Asn Leu Glu Glu Gln Thr Glu Arg Asn Gly Asp Val Ala Asn 385 390 395 400

Leu Glu Asn Glu Ser Lys Val 405

<210> 15

<211> 6

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence:channel protein central pore signature sequence

<400> 15

Thr Val Gly Tyr Gly Asp

1

5

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<210> 16
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<211> 160

<212> PRT

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence:mutated prokaryotic kcsa potassium channel protein of Streptomyces lividans

<400> 16

Met Pro Pro Met Leu Ser Gly Leu Leu Ala Arg Leu Val Lys Leu Leu
1 5 10 15

Leu Gly Arg His Gly Ser Ala Leu His Trp Arg Ala Ala Gly Ala Ala 20 25 30

Thr Val Leu Leu Val Ile Val Leu Leu Ala Gly Ser Tyr Leu Ala Val
35 40 45

Leu Ala Glu Arg Gly Ala Pro Gly Ala Ala Leu Ile Ser Tyr Pro Asp 50 55 60

Ala Leu Trp Trp Ser Val Glu Thr Ala Thr Thr Val Gly Tyr Gly Asp
65 70 75 80

Leu Tyr Pro Val Thr Leu Trp Gly Arg Leu Val Ala Val Val Met
85 90 95

Val Ala Gly Ile Thr Ser Phe Gly Leu Val Thr Ala Ala Leu Ala Thr
100 105 110

Trp Phe Val Gly Arg Glu Gln Glu Arg Arg Gly His Phe Val Arg His
115 120 125

Ser Glu Lys Ala Ala Glu Glu Ala Tyr Thr Arg Thr Thr Arg Ala Leu 130 135 140

His Glu Arg Phe Asp Arg Leu Glu Arg Met Leu Asp Asp Asn Arg Arg 145 150 155 160

<210> 17

<211> 1161

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence:nuclec acid encoding mutated prokaryotic kcsa potassium channel protein of Streptomyces lividans

<400> 17

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agegeggege accgggegeg gegetgatet egtateegga egegetgtgg tggteegtgg 540
 agaccgcgac gaccgtcggc tacggcgacc tgtaccccgt gactctgtgg ggccggctcg 600
 tggccgtggt ggtgatggtc gccgggatca cctccttcgg tctggtgacc gccgcgctgg 660
 ccacctggtt cgtcggccgg gaacaagagc gccggggcca cttcgtgcgc cactccgaga 720
 aggeogeoga ggaggogtac acgeggacga ecegggeget gcacgagegt ttegacegtt 780
 tggagcgaat gctcgacgac aaccgccggt gactccgccg gtgaccgccc gagcgaggcc 840
 gcaccgatga gtctgcggcg gttgtgcggt ctacccgtcg acgaagggag cgcaccatgc 900
 gcaagateat catttgcacg tteetgacge tggaeggegt catgcaggeg eegggegee 960
 cggacgagga cgccgagagc ggcttcgaac acggcggctg gcagaagccg gtggacgacg 1020
 acgaggtcgg cacggccatc gccggctggt acgaggactc cgacgccatg ctcctcggcc 1080
 gcaagaccta cgacatcttc gcgtcgtact ggccgaccgc cgaccccgac aacccgttca 1140
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 <210> 18
 <211> 261
 <212> PRT
 <213> Homo sapiens
 <400> 18
 Ile Tyr Tyr Ala Ile Ser Lys Ser Ile Gly Phe Gly Val Asp Thr Trp
 Val Tyr Pro Asn Ile Thr Asp Pro Glu Tyr Gly Tyr Leu Ala Arg Glu
              20
 Tyr Ile Tyr Cys Leu Tyr Trp Ser Thr Leu Thr Leu Thr Thr Ile Gly
 Glu Thr Pro Pro Pro Val Lys Asp Glu Glu Tyr Leu Phe Val Ile Phe
                          55
 Asp Phe Leu Ile Gly Val Leu Ile Phe Ala Thr Ile Val Gly Asn Val
                      70
 Gly Ser Met Ile Ser Asn Met Asn Ala Thr Arg Ala Glu Phe Gln Ala
 Lys Ile Asp Ala Val Lys His Tyr Met Gln Phe Arg Lys Val Ser Lys
             100
                                 105
 Gly Met Glu Ala Lys Val Ile Arg Trp Phe Asp Tyr Leu Trp Thr Asn
                             120
 Lys Lys Thr Val Asp Glu Arg Glu Ile Leu Lys Asn Leu Pro Ala Lys
                         135
 Leu Arg Ala Glu Ile Ala Ile Asn Val His Leu Ser Thr Leu Lys Lys
 145
                     150
                                         155
 Val Arg Ile Phe His Asp Cys Glu Ala Gly Leu Leu Val Glu Leu Val
                 165
                                     170
'Leu Lys Leu Arg Pro Gln Val Phe Ser Pro Gly Asp Tyr Ile Cys Arg
             180
                                 185
 Lys Gly Asp Ile Gly Lys Glu Met Tyr Ile Ile Lys Glu Gly Lys Leu
                             200
```

Ala Val Val Ala Asp Asp Gly Val Thr Gln Tyr Ala Leu Leu Ser Ala

215

210

Gly Ser Cys Phe Gly Glu le Ser Ile Leu Asn Ile Lys Gly Ser Lys 225 230 235 240

Met Gly Asn Arg Arg Thr Ala Asn Ile Arg Ser Leu Gly Tyr Ser Asp 245 250 255

Leu Phe Cys Leu Ser 260

<210> 19

<211> 690

<212> PRT

<213> Homo sapiens

<400> 19

Met Lys Leu Ser Met Lys Asn Asn Ile Ile Asn Thr Gln Gln Ser Phe 1 5 10 15

Val Thr Met Pro Asn Val Ile Val Pro Asp Ile Glu Lys Glu Ile Arg
20 25 30

Arg Met Glu Asn Gly Ala Cys Ser Ser Phe Ser Glu Asp Asp Asp Ser 35 40 45

Ala Tyr Thr Ser Glu Glu Ser Glu Asn Glu Asn Pro His Ala Arg Gly
50 55 60

Ser Phe Ser Tyr Lys Ser Leu Arg Lys Gly Gly Pro Ser Gln Arg Glu 65 70 75 80

Gln Tyr Leu Pro Gly Ala Ile Ala Ile Phe Asn Val Asn Asn Ser Ser 85 90 95

Asn Lys Asp Gln Glu Pro Glu Glu Lys Lys Lys Lys Lys Glu Lys
100 105 110

Lys Ser Lys Ser Asp Asp Lys Asn Glu Asn Lys Asn Asp Pro Glu Lys
115 120 125

Lys Lys Lys Lys Lys Asp Lys Glu Lys Lys Lys Glu Glu Lys Ser

Lys Asp Lys Lys Glu His His Lys Lys Glu Val Val Val Ile Asp Pro 145 150 155 160

Ser Gly Asn Thr Tyr Tyr Asn Trp Leu Phe Cys Ile Thr Leu Pro Val 165 170 175

Gln Ser Asp Tyr Leu Glu Tyr Trp Leu Ile Leu Asp Tyr Val Ser Asp 195 200 205

Ile Val Tyr Leu Ile Asp Met Phe Val Arg Thr Arg Thr Gly Tyr Leu 210 215 220

Glu Gln Gly Leu Leu Val Lys Glu Glu Leu Lys Leu Ile Asn Lys Tyr 225 230 235 240

- Lys Ser Asn Leu Gln Phe Lys Leu Asp Val Leu Ser Leu Ile Pro Thr 245 250 255
- Asp Leu Leu Tyr Phe Lys Leu Gly Trp Asn Tyr Pro Glu Ile Arg Leu 260 265 270
- Asn Arg Leu Leu Arg Phe Ser Arg Met Phe Glu Phe Phe Gln Arg Thr 275 280 285
- Glu Thr Arg Thr Asn Tyr Pro Asn Ile Phe Arg Ile Ser Asn Leu Val 290 295 300
- Met Tyr Ile Val Ile Ile Ile His Trp Asn Ala Cys Val Phe Tyr Ser 305 310 315 320
- Ile Ser Lys Ala Ile Gly Phe Gly Asn Asp Thr Trp Val Tyr Pro Asp 325 330 335
- Ile Asn Asp Pro Glu Phe Gly Arg Leu Ala Arg Lys Tyr Val Tyr Ser 340 345 350
- Leu Tyr Trp Ser Thr Leu Thr Leu Thr Thr Ile Gly Glu Thr Pro Pro 355 360 365
- Pro Val Arg Asp Ser Glu Tyr Val Phe Val Val Val Asp Phe Leu Ile 370 375 380
- Gly Val Leu Ile Phe Ala Thr Ile Val Gly Asn Ile Gly Ser Met Ile 385 390 395 400
- Ser Asn Met Asn Ala Ala Arg Ala Glu Phe Gln Ala Arg Ile Asp Ala 405 410 415
- Ile Lys Gln Tyr Met His Phe Arg Asn Val Ser Lys Asp Met Glu Lys
 420 425 430
- Arg Val Ile Lys Trp Phe Asp Tyr Leu Trp Thr Asn Lys Lys Thr Val
 435
 440
 445
- Asp Glu Lys Glu Val Leu Lys Tyr Leu Pro Asp Lys Leu Arg Ala Glu 450 455 460
- Ile Ala Ile Asn Val His Leu Asp Thr Leu Lys Lys Val Arg Ile Phe 465 470 475 480
- Ala Asp Cys Glu Ala Gly Leu Leu Val Glu Leu Val Leu Lys Leu Gln
 485 490 495
- Pro Gln Val Tyr Ser Pro Gly Asp Tyr Ile Cys Lys Lys Gly Asp Ile
 500 505 . 510
- Gly Arg Glu Met Tyr Ile Ile Lys Glu Gly Lys Leu Ala Val Val Ala 515 520 525
- Asp Asp Gly Val Thr Gln Phe Val Val Leu Ser Asp Gly Ser Thr Phe 530 540
- Gly Glu Ile Ser Ile Leu Asn Ile Lys Gly Ser Lys Ala Gly Asn Arg 545 550 555 560

Arg Thr Ala Asn Ile Lys Ser Ile Gly Tyr Ser Asp Leu Phe Cys Leu 565 570 575

Ser Lys Asp Asp Leu Met Glu Ala Leu Thr Glu Tyr Pro Asp Ala Lys 580 585 590

Thr Met Leu Glu Glu Lys Gly Lys Gln Ile Leu Met Lys Asp Gly Leu
595 600 605

Leu Asp Leu Asnalle Ala Asn Ala Gly Ser Asp Pro Lys Asp Leu Glu 610 620

Glu Lys Val Thr Arg Met Glu Gly Ser Val Asp Leu Leu Gln Thr Arg 625 630 635

Phe Ala Arg Ile Leu Ala Glu Tyr Glu Ser Met Gln Gln Lys Leu Lys 645 650 655

Gln Arg Leu Thr Lys Val Glu Lys Phe Leu Lys Pro Leu Ile Asp Thr
660 665 670

Glu Phe Ser Ser Ile Glu Gly Pro Trp Ser Glu Ser Gly Pro Ile Asp 675 680 685

Ser Thr 690

<210> 20

<211> 40

<212> PRT

<213> Drosophila melanogaster

<400× 20

Ala Glu Ala Gly Ser Glu Asn Ser Phe Phe Lys Ser Ile Pro Asp Ala
1 5 10 15

Phe Trp Trp Ala Val Val Thr Met Thr Thr Val Gly Tyr Gly Asp Met 20 25 30

Thr Pro Val Gly Phe Trp Gly Lys

<210> 21

<211> 40

<212> PRT

<213> Rattus norvegicus

<400> 21

Ala Asn His Thr Pro Cys Val Glu Asn Ile Asn Gly Met Thr Ser Ala 1 5 10 15

Phe Leu Phe Ser Leu Glu Thr Gln Val Thr Ile Gly Tyr Gly Phe Arg 20 25 30

Cys Val Thr Glu Gln Cys Ala Thr 35 40

```
<210> 22
<211> 42
<212> PRT
<213> Methanococcus jannaschii
Glu Ser Val Ile Leu Met Thr Val Glu Gly Trp Asp Phe Phe Thr Ala
                                      10
Phe Tyr Thr Ala Val Val Thr Ile Ser Thr Val Gly Tyr Gly Asp Tyr
                                  25
Thr Pro Gln Thr Phe Leu Gly Lys Leu Ser
      . 35
<210> 23
<211> 40
<212> PRT
<213> Streptomyces lividans
<400> 23
Val Leu Ala Glu Arg Pro Gly Ala Gln Leu Ile Thr Tyr Pro Arg Ala
Leu Trp Trp Ser Val Glu Thr Ala Thr Thr Val Gly Tyr Gly Asp Leu
Tyr Pro Val Thr Leu Trp Gly Arg
         35 .
<210> 24
<211> 5
<212> PRT
<213> Artificial Sequence
<223> Description of Artificial Sequence: channel protein
      central pore signature sequence
<400> 24
Thr Val Gly Tyr Gly
  1
<210> 25
<211> 58
<212> PRT
<213> Streptomyces lividans
Thr Tyr Pro Arg Ala Leu Trp Trp Ser Val Glu Thr Ala Thr Thr Val
Gly Tyr Gly Asp Leu Tyr Pro Val Thr Leu Trp Gly Arg Leu Val Ala
             20
                                 25
                                                      3.0
```

Val Val Met Val Ala Gly Ile Thr Ser Phe Gly Leu Val Thr Ala

40

35

```
Ala Leu Ala Thr Trp Phe Val Gly Arg Glu 50 55
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<210> 26

<211> 58

<212> PRT

<213> Escherichia coli

<400> 26

Ser Leu Met Thr Ala Phe Tyr Phe Ser Ile Glu Thr Met Ser Thr Val 1 5 10 15

Gly Tyr Gly Asp Ile Val Pro Val Ser Glu Ser Ala Arg Leu Phe Thr

Ile Ser Val Ile Ile Ser Gly Ile Thr Val Phe Ala Thr Ser Met Thr 35 40 45

Ser Ile Phe Gly Pro Leu Ile Arg Gly Gly 50 55

<210> 27

<211> 58

<212> PRT

<213> Drosophila melanogaster

<400> 27

Ser Ile Pro Asp Ala Phe Trp Trp Ala Val Val Thr Met Thr Thr Val 1 5 10 15

Ser Leu Cys Val Ile Ala Gly Val Leu Thr Ile Ala Leu Pro Val Pro 35 40 45

Val Ile Val Ser Asn Phe Asn Tyr Phe Tyr
50 55

<210> 28

<211> 58

<212> PRT

<213> Homo sapiens

<400> 28

Ser Ile Pro Asp Ala Phe Trp Trp Ala Val Val Ser Met Thr Thr Val

1 10 15

Gly Tyr Gly Asp Met Tyr Pro Val Thr Ile Gly Gly Lys Ile Val Gly 20 25 30

Ser Leu Cys Ala Ile Ala Gly Val Leu Thr Ile Ala Leu Pro Val Pro
35 40 45

Val Ile Val Ser Asn Phe Asn Tyr Phe Tyr

```
<210> 29
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<211> 58

<212> PRT

<213> Homo sapiens

<400> 29

Ser Ile Pro Ala Ser Phe Trp Trp Ala Thr Ile Thr Met Thr Thr Val 1 5 10 15

Gly Tyr Gly Asp Ile Tyr Pro Lys Thr Leu Leu Gly Lys Ile Val Gly
20 25 30

Gly Leu Cys Cys Ile Ala Gly Val Leu Val Ile Ala Leu Pro Ile Pro 35 40 45

Ile Ile Val Asn Asn Phe Ser Glu Phe Tyr
50 55

<210> 30

<211> 58

<212> PRT

<213> Paramecium tetraurelia

<400> 30

Gln Tyr Leu His Ser Leu Tyr Trp Ser Ile Ile Thr Met Thr Thr Ile 1 5 10 15

Gly Tyr Gly Asp Ile Thr Pro Gln Asn Leu Arg Glu Arg Val Phe Ala
20 25 30

Val Gly Met Ala Leu Ser Ala Val Gly Val Phe Gly Tyr Ser Ile Gly 35 40 45

Asn Ile Asn Ser Ile Tyr Ala Glu Trp Ser 50 55

<210> 31

<211> 58

<212> PRT

<213> Caenorhabditis elegans

<400> 31

Ser Ile Pro Leu Gly Leu Trp Trp Ala Ile Cys Thr Met Thr Thr Val 1 5 10 15

Gly Tyr Gly Asp Met Thr Pro His Thr Ser Phe Gly Arg Leu Val Gly
20 25 30

Ser Leu Cys Ala Val Met Gly Val Leu Thr Ile Ala Leu Pro Val Pro 35 40 45

Val Ile Val Ser Asn Phe Ala Met Phe Tyr

<210> 32

<211> 58

<212> PRT

<213> Mus musculus

```
<400> 32
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Thr Tyr Trp Glu Cys Val Tyr Leu Leu Met Val Thr Met Ser Thr Val 1 5 10 15

Gly Tyr Gly Asp Val Tyr Ala Lys Thr Thr Leu Gly Arg Leu Phe Met 20 25 30

Val Phe Phe Ile Leu Gly Gly Leu Ala Met Phe Ala Ser Tyr Val Pro 35 40 45

Glu Ile Ile Glu Leu Ile Gly Asn Arg Lys
50 55

<210> 33

<211> 58

<212> PRT

<213> Homo sapiens

<400> 33

Asn Phe Leu Gly Ala Met Trp Leu Ile Ser Ile Thr Phe Leu Ser Ile 1 5 10 15

Gly Tyr Gly Asp Met Val Pro His Thr Tyr Cys Gly Lys Gly Val Cys
20 25 30

Leu Leu Thr Gly Ile Met Gly Ala Gly Cys Thr Ala Leu Val Val Ala
35 40 45

Val Val Ala Arg Lys Leu Glu Leu Thr Lys
50 55

<210> 34

<211> 58

<212> PRT

<213> Arabidopsis thaliana

<400> 34

Arg Tyr Val Thr Ser Met Tyr Trp Ser Ile Thr Thr Leu Thr Thr Val 1 5 10 15

Gly Tyr Gly Asp Leu His Pro Val Asn Thr Lys Glu Met Ile Phe Asp

Ile Phe Tyr Met Leu Phe Asn Leu Gly Leu Thr Ala Tyr Leu Ile Gly
35 40 45

Asn Met Thr Asn Leu Val Val His Gly Thr 50 55

<210> 35

<211> 58

<212> PRT

<213> Homo sapiens

<400> 35

Lys Tyr Val Thr Ala Leu Tyr Phe Thr Phe Ser Ser Leu Thr Ser Val 1 5 10 15 Gly Phe Gly Asn Val Ser Pro Asn Thr Asn Ser Glu Lys Ile Phe Ser 20 25 30

Ile Cys Val Met Leu Ile Gly Ser Leu Met Tyr Ala Ser Ile Phe Gly 35 ' 40 45

Asn Val Ser Ala Ile Ile Gln Arg Leu Tyr 50 55

<210> 36

<211> 60

<212> PRT

<213> Rattus norvegicus

<400> 36

Gly Met Thr Ser Ala Phe Leu Phe Ser Leu Glu Thr Gln Val Thr Ile 1 5 10 15

Gly Tyr Gly Phe Arg Phe Val Thr Glu Gln Cys Ala Thr Ala Ile Phe 20 25 30

Leu Leu Ile Phe Gln Ser Ile Leu Gly Val Ile Ile Asn Ser Phe Met
35 40 45

Cys Gly Ala Ile Leu Ala Lys Ile Ser Arg Pro Lys 50 55 60

<210> 37

<211> 60

<212> PRT

<213> Homo sapiens

<400> 37

Gly Phe Val Ser Ala Phe Leu Phe Ser Ile Glu Thr Glu Thr Thr Ile

1 5 10 15

Gly Tyr Gly Tyr Arg Val Ile Thr Asp Lys Cys Pro Glu Gly Ile Ile 20 25 30

Leu Leu Ile Gln Ser Val Leu Gly Ser Ile Val Asn Ala Phe Met
35 40 45

Val Gly Cys Met Phe Val Lys Ile Ser Gln Pro Lys 50 55 60

<210> 38

<211> 57

<212> PRT

<213> Homo sapiens

<400> 38

Glu Tyr Ile Tyr Cys Leu Tyr Trp Ser Thr Leu Thr Leu Thr Thr Ile

1 10 15

Gly Glu Thr Pro Pro Pro Val Lys Asp Glu Glu Tyr Leu Phe Val Ile 20 25 30

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Phe Asp Phe Leu Ile Gly Val Leu Ile Phe Ala Thr Ile Val Gly Asn
                             40
Val Gly Ser Met Ile Ser Asn Met Asn
     50
<210> 39
<211> 57
<212> PRT
<213> Homo sapiens
Lys Tyr Val Tyr Ser Leu Tyr Trp Ser Thr Leu Thr Leu Thr Thr Ile
Gly Glu Thr Pro Pro Pro Val Arg Asp Ser Glu Tyr Val Phe Val Val
Val Asp Phe Leu Ile Gly Val Leu Ile Phe Ala Thr Ile Val Gly Asn
                             40
Ile Gly Ser Met Ile Ser Asn Met Asn
<210> 40
<211> 4
<212> PRT
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence:selectivity
     filter sequence
<400> 40
Val Gly Tyr Gly
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<210> 41
<211> 36
<212> PRT
<213> Streptomyces lividans
<400> 41
Glu Arg Gly Ala Pro Gly Ala Gln Leu Ile Thr Tyr Pro Arg Ala Leu
Trp Trp Ser Val Glu Thr Ala Thr Thr Val Gly Tyr Gly Asp Leu Tyr
                                 25
Pro Val Thr Leu
        35
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<210> 42 <211> 36

<212> PRT

<213> Drosophila melanogaster

Trp Trp Ala Val Val Thr Met Thr Thr Val Gly Tyr Gly Asp Met Thr 20 25 30

Pro Val Gly Phe 35